FIGURE 1
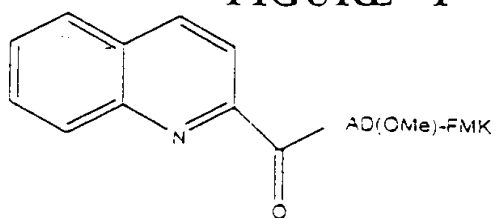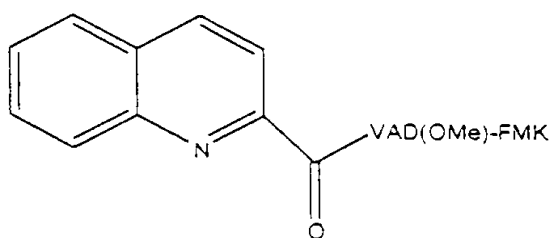

Quin-AD(OMe)-FMK M.Wt:389

FIGURE 1A

Quin-VAD(OMe)-FMK
M.Wt:488; C24H19N4O6F

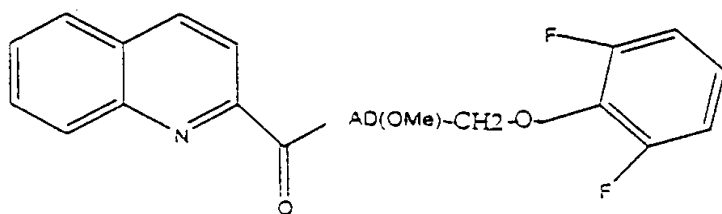# FIGURE 2

FIGURE 2A
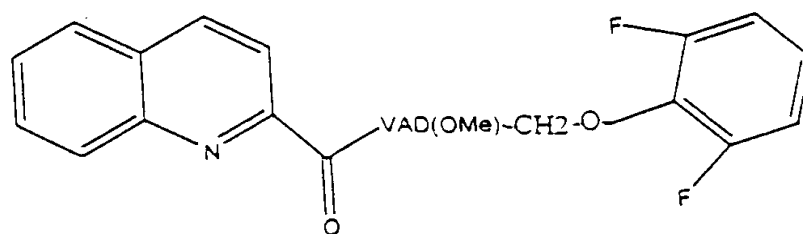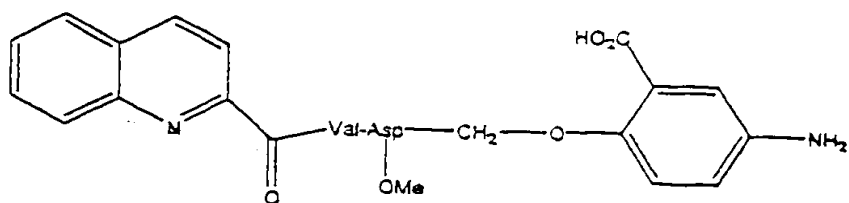
FIGURE 3

FIGURE 4
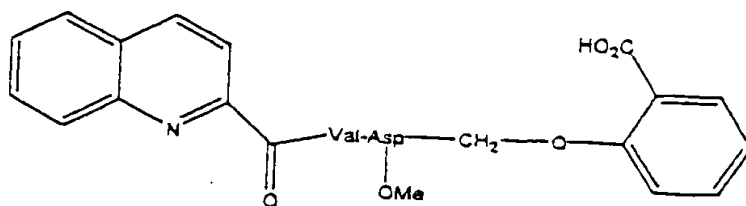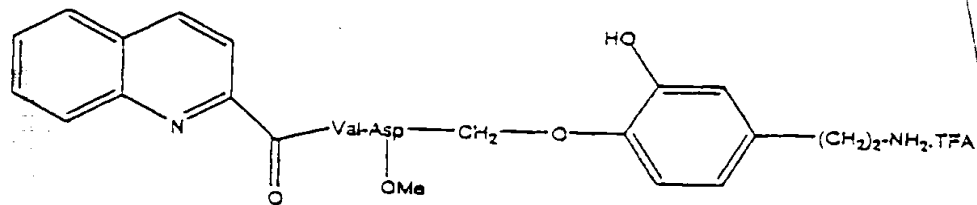

FIGURE 5
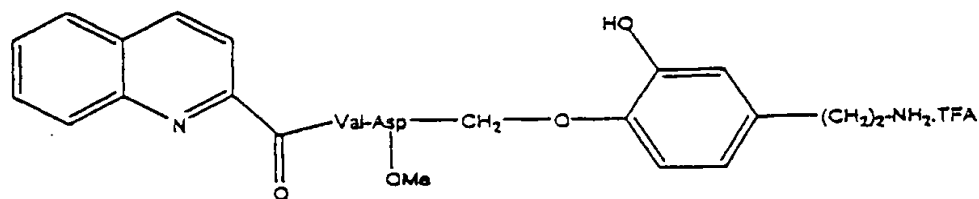
FIGURE 6
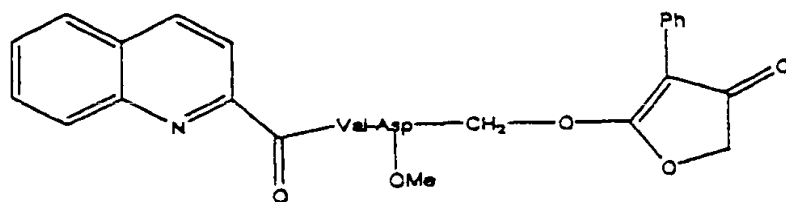
FIGURE 7

FIGURE 8

Caspase 9

inh conc	log of con	% inhib
0.005uM	-2.301	0
0.01uM	-2	0
.025uM	-1.602	0
.05uM	-1.301	0
.1uM	-1	0
0.5uM	-0.301	0
1uM	0	16.2
2.5uM	0.3979	21.8
5uM	0.6989	47.4
10uM	1	62
25uM	1.398	82.4
50uM	1.6989	92.6

Q-(C=O)-VD(OMe)-CH$_2$-ASA
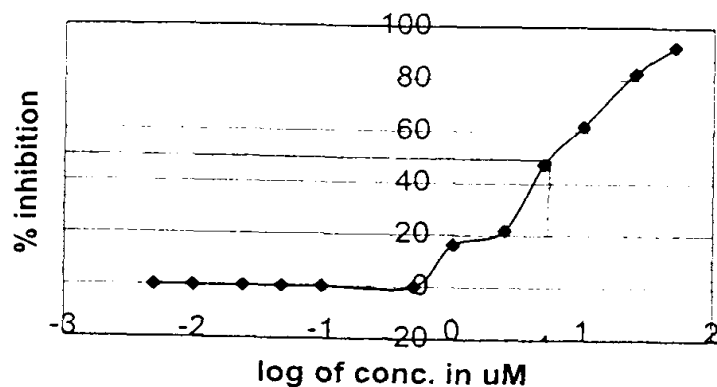
FIGURE 9

Caspase 8

inh conc	log of con	% inhib
0.005uM	-2.301	0
0.01uM	-2	0
.025uM	-1.602	0
.05uM	-1.301	0
.1uM	-1	0
0.5uM	-0.301	4.7
1uM	0	5.5
2.5uM	0.3979	21.1
5uM	0.6989	45.5
10uM	1	73.6
25uM	1.398	96.8
50uM	1.6989	99.8

Q-(C=O)-VD(OMe)-CH$_2$-ASA
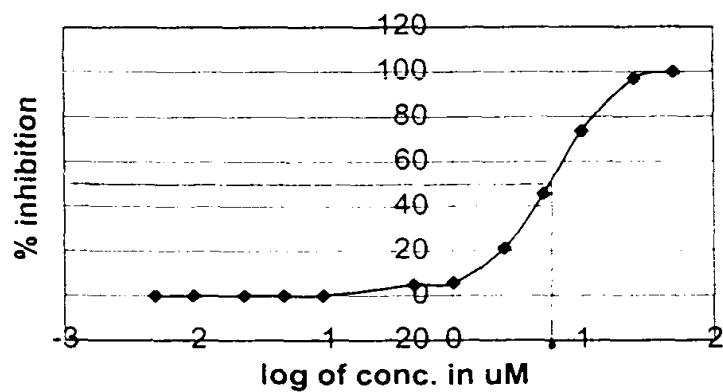
FIGURE 10

Caspase 1

inh conc	log of con	% inhib
.025uM	-1.602	0
.05uM	-1.301	0
.1uM	-1	0
0.5uM	-0.301	18.2
1uM	0	34.8
2.5uM	0.3979	69.7
5uM	0.6989	100
10uM	1	100
25uM	1.398	100
50uM	1.6989	100
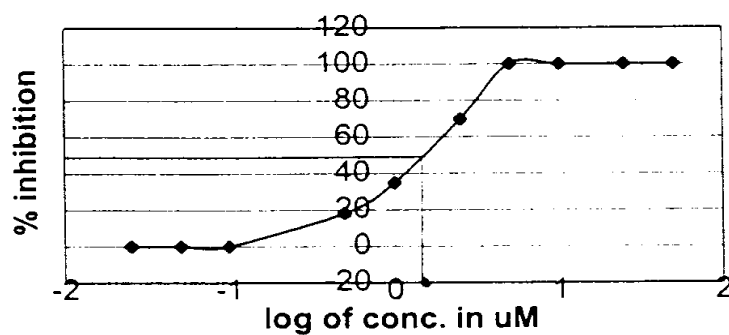
Q-(C=O)-VD(OMe)-CH$_2$-ASA

FIGURE 11

Caspase 3

inh conc	log of con	% inhib
0.005uM	-2.301	0
0.01uM	-2	0
.025uM	-1.602	2.3
.05uM	-1.301	9.1
.1uM	-1	6.4
0.5uM	-0.301	29.3
1uM	0	45
2.5uM	0.3979	74.8
5uM	0.6989	91.5
10uM	1	98.2
25uM	1.398	100
50uM	1.6989	100
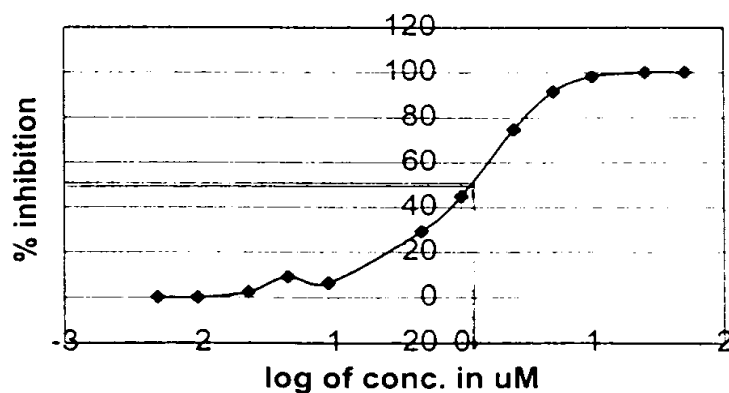
Q-(C=O)-VD(OMe)-CH$_2$-ASA

FIGURE 12

Caspase 1

inh conc	log of con	% inhib
.0025uM	-2.602	3.14
.005uM	-2.301	2.6
.01uM	-2	1.4
.025uM	-1.602	10.3
.05uM	-1.301	8.3
.1uM	-1	23.7
0.5uM	-0.301	50.9
1uM	0	66.29
2.5uM	0.3979	90.3
5uM	0.6989	96.3
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

Indole-(C=O)-VD(OMe)-CH$_2$-OPh
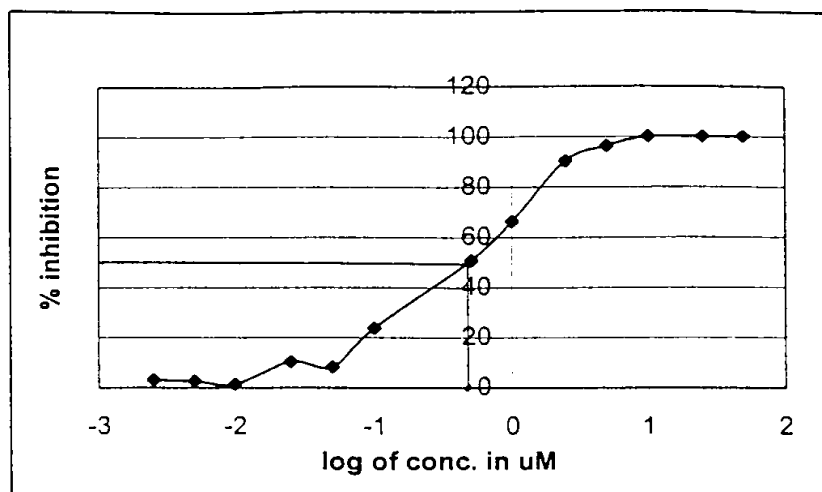
FIGURE 13

Caspase 1

inh conc	log of con	% inhib
.0025uM	-2.602	16 3
.005uM	-2.301	19 4
.01uM	-2	22 6
.025uM	-1.602	42.86
.1uM	-1	44
0.5uM	-0.301	74
1uM	0	87 4
2.5uM	0.3979	97.1
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

Melatonin-VD(OMe)-CH$_2$-OPh
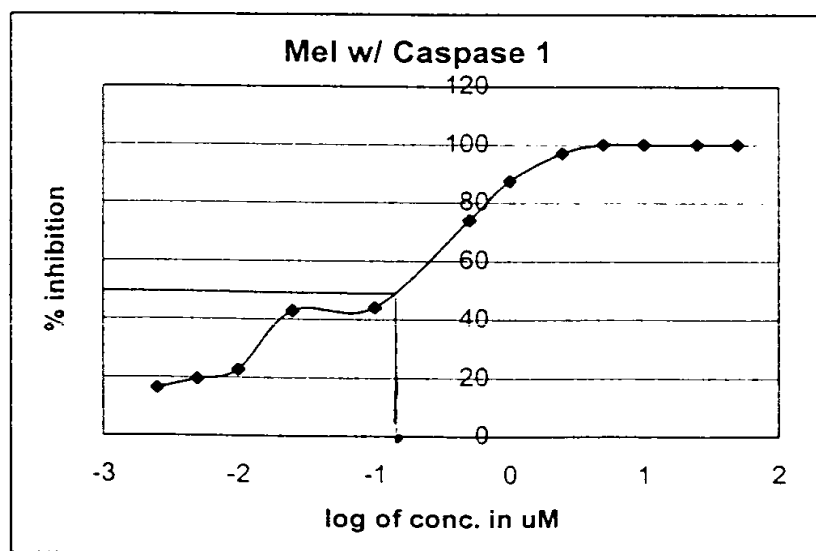
FIGURE 14

Caspase 1

Bzl-Melatonin-VD(OMe)-CH₂-OPh

inh conc	log of con	% inhib
0.0025uM	-2.602	0
0.005uM	-2.301	0
0.01uM	-2	0
.025uM	-1.602	0
.05uM	-1.301	7.3
.1uM	-1	26.8
0.5uM	-0.301	93.4
1uM	0	99.6
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.398	100
50uM	1.6989	100
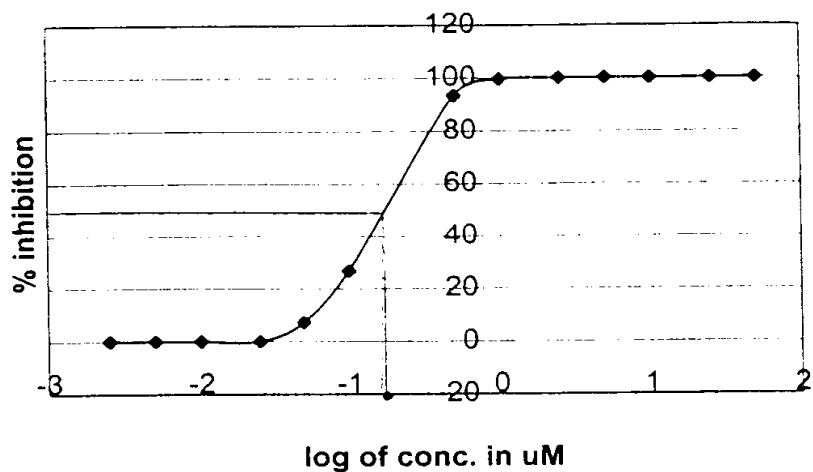
log of conc. in uM

FIGURE 15

Caspase 1

HydroxyTryptophan-VD(OMe)-CH₂-OPh

inh conc	log of con	% inhib
0.0025uM	-2.602	38.4
0.005uM	-2.301	25.7
0.01uM	-2	29.6
.025uM	-1.602	23
.05uM	-1.301	44.3
0.5uM	-0.301	57.2
1uM	0	91.4
2.5uM	0.3979	95
5uM	0.6989	96.9

log of conc. in uM
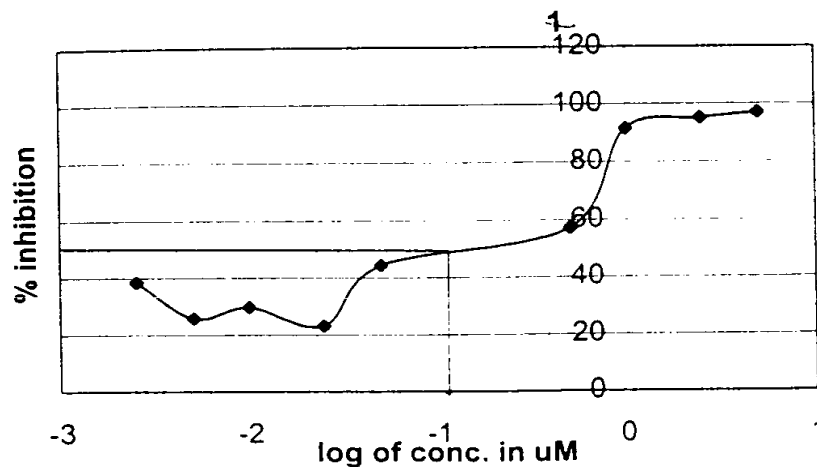
FIGURE 16

Caspase 1

TRP-VD(OCH₃)-CH₂-OPh · TFA

inh conc	log of con	% inhib
0.0025uM	-2.602	0
0.005uM	-2.301	0
0.01uM	-2	0
.025uM	-1.602	0
.05uM	-1.301	0
.1uM	-1	20.7
0.5uM	-0.301	42.7
1uM	0	81.7
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.398	100
50uM	1.6989	100
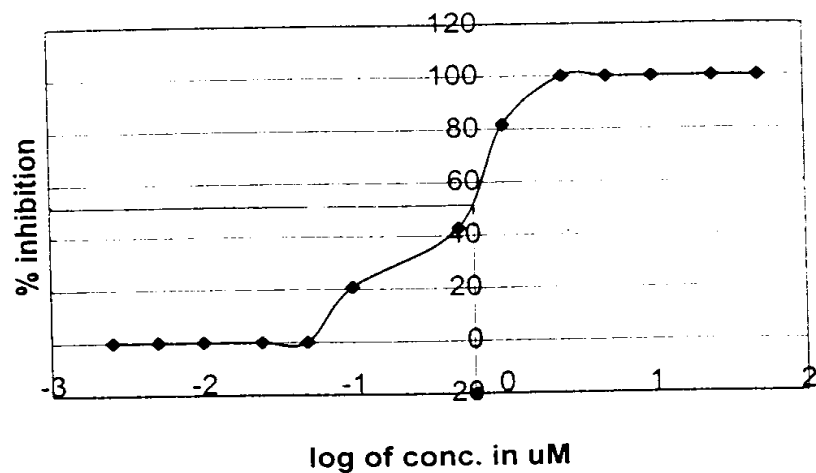
log of conc. in uM

FIGURE 17A

Caspase 9

inh conc	log of con	% inhib
.025uM	-1.602	33.6
.05uM	-1.301	43.9
.1uM	-1	58.7
0 5uM	-0.301	90.7
1uM	0	94.7
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

Q-(C=O)-L-D-(OMe)-CH$_2$-F (the FMK)
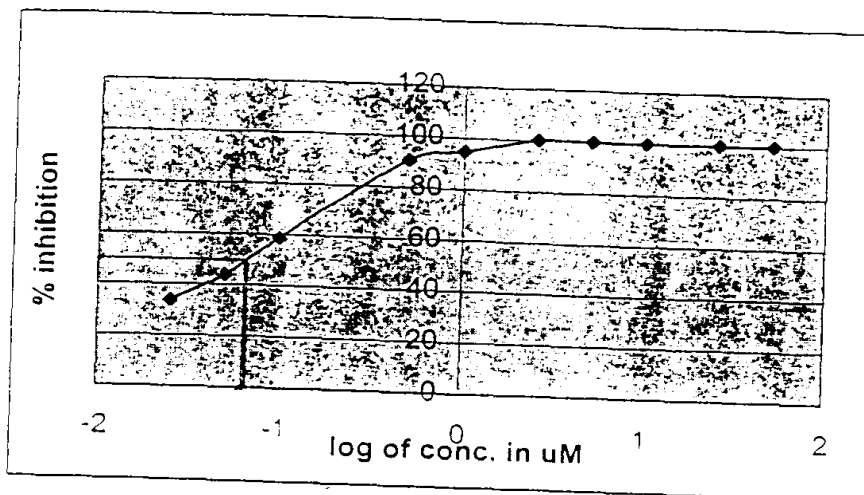
FIGURE 17B

Caspase 9

inh conc	log of con	% inhib
.025uM	-1.602	25.7
.05uM	-1.301	37.3
.1uM	-1	58.9
0 5uM	-0.301	88.9
1uM	0	94.9
2.5uM	0.3979	96.1
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

Q-(C=O)-L-D-(OMe)-CH$_2$-F (the FMK)
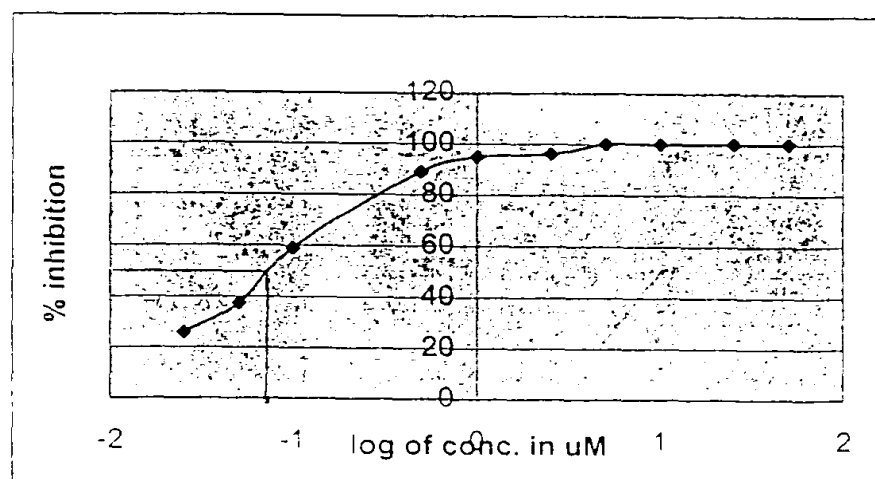
FIGURE 18A

Caspase 9

Q-(C=O)-V-D-(OCH₃)-CH₂-F (the FMK)

nn conc	log of con	% inhib
.025uM	-1.602	47.3
.05uM	-1.301	64.4
.1uM	-1	81.2
0.5uM	-0.301	97.8
1uM	0	99.5
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100
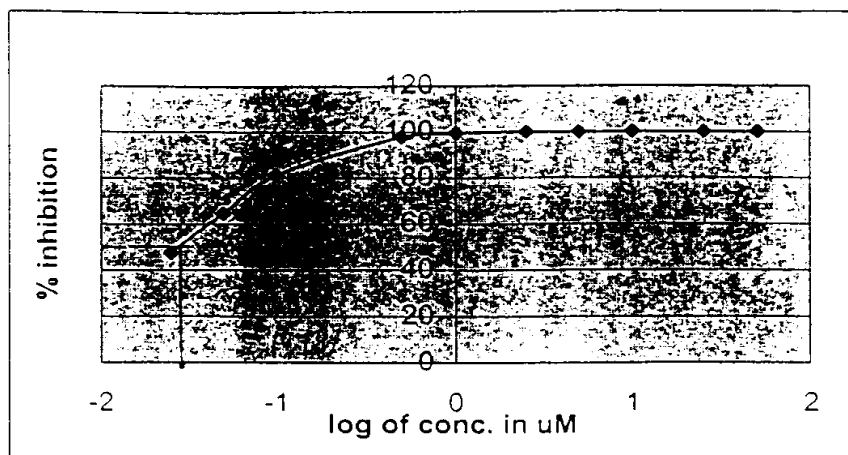
FIGURE 18B

Caspase 9

Q-(C=O)-V-D-(OCH₃)-CH₂-F (the FMK)

nn conc	log of con	% inhib
.025uM	-1.602	62.2
.05uM	-1.301	76.3
.1uM	-1	81.3
0.5uM	-0.301	99.1
1uM	0	100
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100
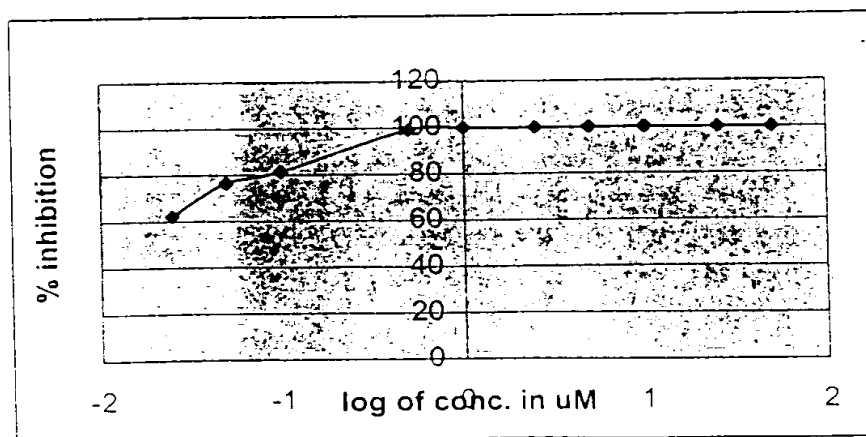
FIGURE 19

Caspase 1

inh conc	log of con	% inhib
0.0025uM	-2.602	0
0.005uM	-2.301	3.2
0.01uM	-2	0
.025uM	-1.602	1.1
.05uM	-1.301	0
.1uM	-1	0
0.5uM	-0.301	15.3
1uM	0	33.7
2.5uM	0.3979	72.1
5uM	0.6989	97.4
10uM	1	100
25uM	1.398	100
50uM	1.6989	100

Q-(C=O)-VD(OMe)-CH$_2$-SA
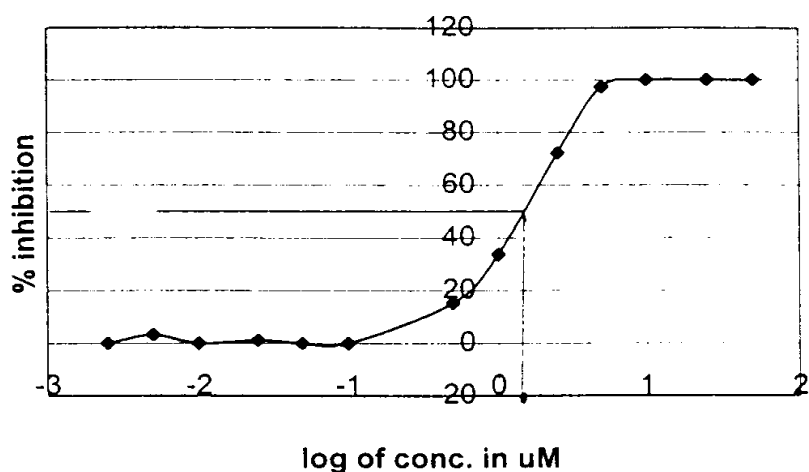
log of conc. in uM

FIGURE 20

Caspase 3

inh conc	log of con	% inhib
0.005uM	-2.301	0
0.01uM	-2	0
.025uM	-1.602	0.57
.05uM	-1.301	2.8
.1uM	-1	18.3
0.5uM	-0.301	32.4
1uM	0	54.7
2 5uM	0.3979	87.8
5uM	0.6989	97.6
10uM	1	99.7
25uM	1.398	100
50uM	1.6989	100

Q-(C=O)-VD(OMe)-CH$_2$-SA
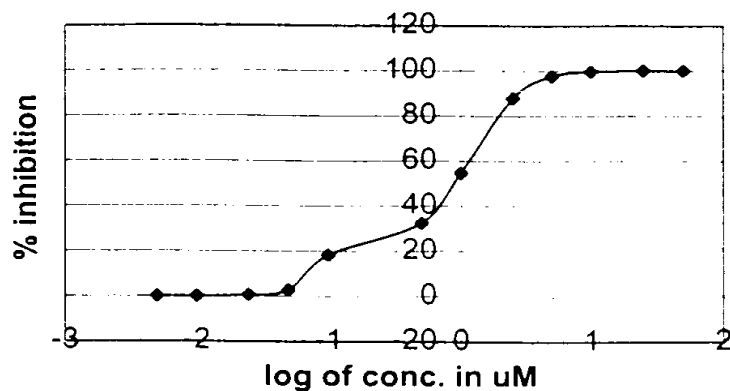
log of conc. in uM

FIGURE 21

Q-(C=O)-L-D-CH₂-OPh

Caspase 1

inh conc	log of con	% inhib
.025uM	-1.602	19
.05uM	-1.301	22
.1uM	-1	19
0 5uM	-0.301	46.7
1uM	0	69.5
2 5uM	0.3979	92.7
5uM	0.6989	98.5
10uM	1	87.3
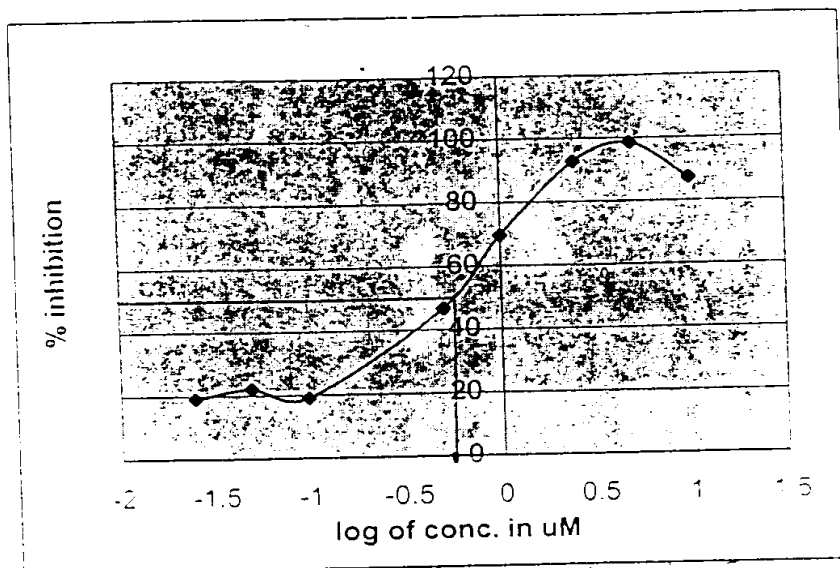
FIGURE 22

Q-(C=O)-V-D-CH₂-OPh

Caspase 1

inh conc	log of con	% inhib
.025uM	-1.602	39.8
.05uM	-1.301	55.98
.1uM	-1	67.2
0 5uM	-0.301	95.8
1uM	0	98.5
2 5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
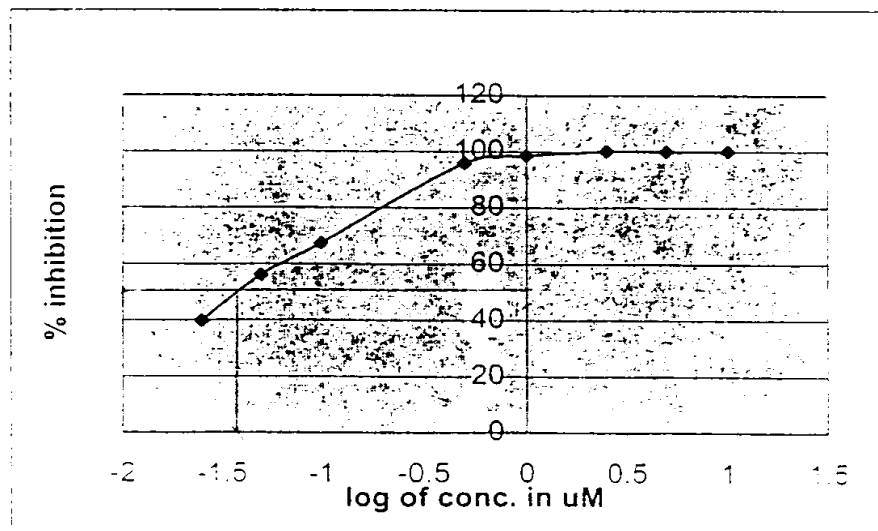
FIGURE 25A

Non esterase treated Inhibitor D with Caspase 3

$$Q\text{-}(C=O)\text{-}L\text{-}D\text{-}(OMe)\text{-}CH_2\text{-}F$$
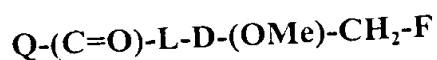
inh conc	log of con	% inhib
0.25uM	-1.602	37.8
0.5uM	-1.301	52
1uM	-1	73
0.5uM	-0.301	100
1uM	0	100
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

Non est trtd Id with Casp 3
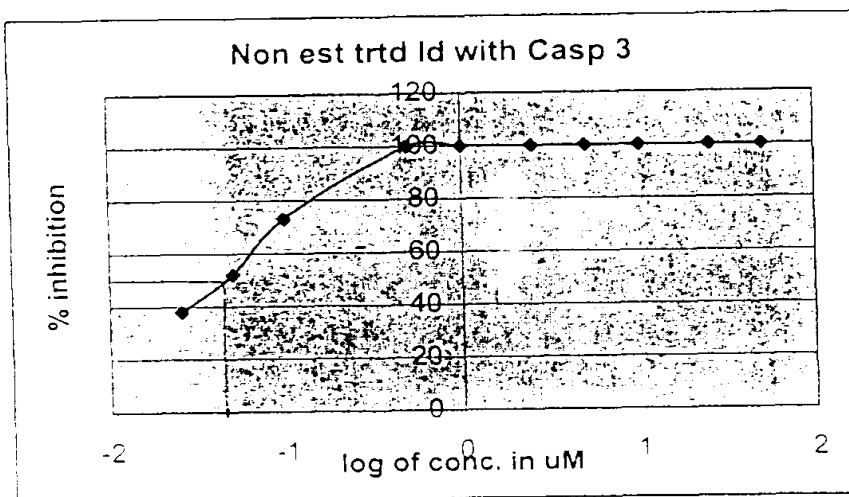
FIGURE 25B

Esterase treated Inhibitor D with Caspase 3

$$Q\text{-}(C=O)\text{-}L\text{-}D\text{-}(OMe)\text{-}CH_2\text{-}F$$
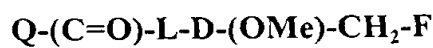
inh conc	log of con	% inhib
0.25uM	-1.602	38.2
0.5uM	-1.301	68.9
1uM	-1	80.7
0.5uM	-0.301	97.6
1uM	0	96.6
2.5uM	0.3979	96.2
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

Est trtd Id w/ Casp 3
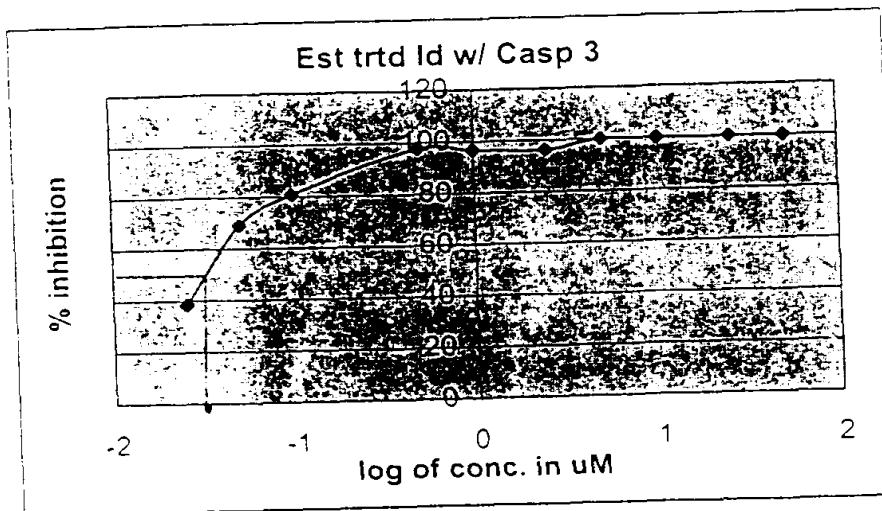
FIGURE 23

Esterase treated Inhibitor C with Caspase 1
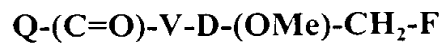
inh conc	log of con	% inhib
.025uM	-1.602	40.1
.05uM	-1.301	54.9
.1uM	-1	73.2
0.5uM	-0.301	81.7
1uM	0	100
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

$Q\text{-}(C\text{=}O)\text{-}V\text{-}D\text{-}(OMe)\text{-}CH_2\text{-}F$

Est trtd Ic w/ Caspase 1
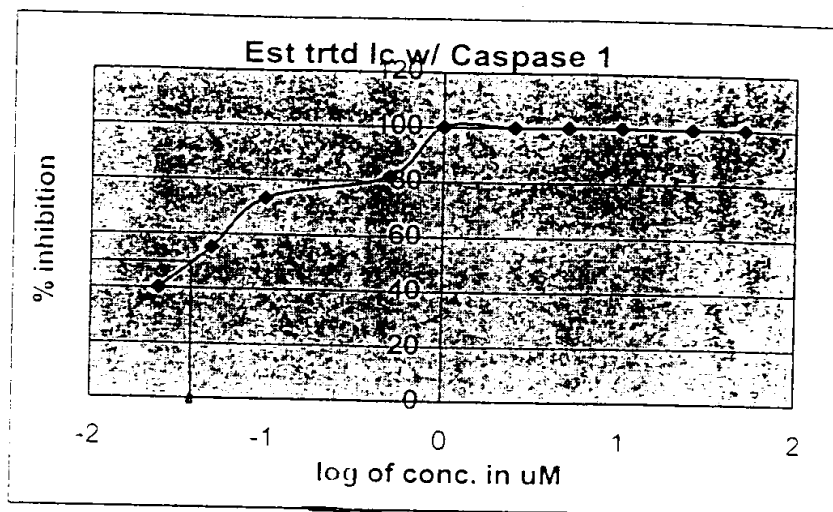
FIGURE 24

Esterase treated Inhibitor D with Casp 1
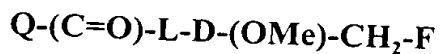
inh conc	log of con	% inhib
.025uM	-1.602	0
.05uM	-1.301	33.8
.1uM	-1	63.4
0.5uM	-0.301	85.2
2.5uM	0.3979	95.2
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.6979	100

$Q\text{-}(C\text{=}O)\text{-}L\text{-}D\text{-}(OMe)\text{-}CH_2\text{-}F$

est trtd Id w/ Casp 1

FIGURE 26

Q-LD-OPh

0.05 uM	-1.301	5.5
0.1 uM	-1	11
0.5 uM	-0.301	46
1 uM	0	68
2.5 uM	0.3979	86.8
5 uM	0.6989	94.5
0 uM	1	100
25 uM	1.3979	100
50 uM	1.6989	100

le inhibition of Caspase 1

FIGURE 27

Q-VD-OPh

It [conc.] log conc.		% inhibition
0.05 uM	-1.301	51.6
0.1 uM	-1	75
0.5 uM	-0.301	97.8
1 uM	0	96.7
2.5 uM	0.3979	98.9
5 uM	0.6989	98.9
uM	1	100
25 uM	1.3979	100
50 uM	1.6989	100

If inhibition of Caspase 1
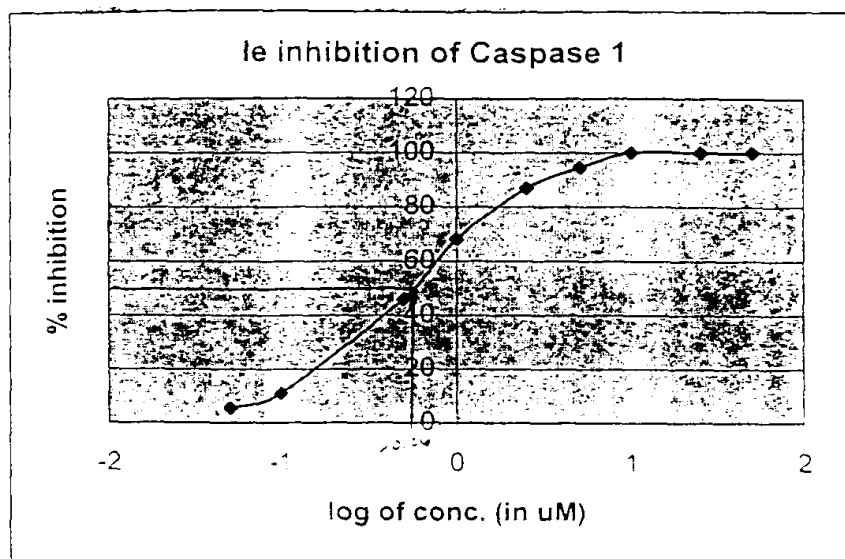
Caspase 3 w/ IE

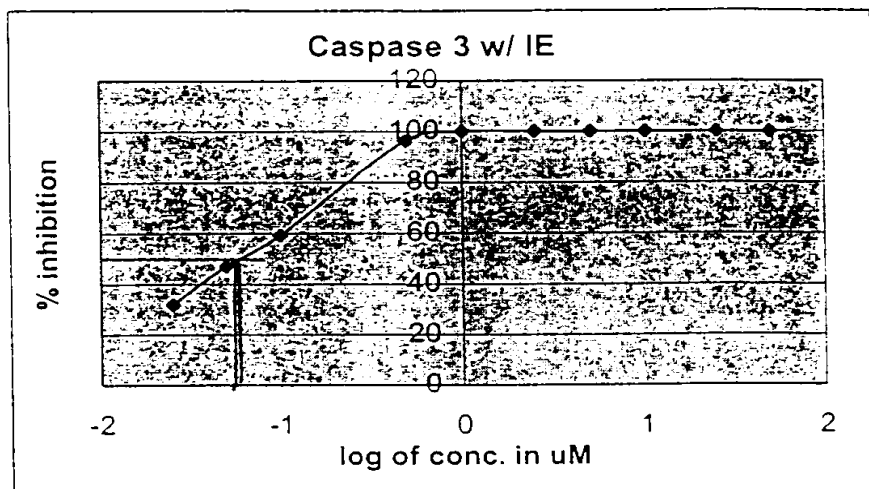| inh conc	log of con	% inhib
.025uM	-1.602	31.85
.05uM	-1.301	47.1
.1uM	-1	59.2
0.5uM	-0.301	96.2
1uM	0	100
2.5uM	0.3979	100
5uM	0.6989	100
10uM	1	100
25uM	1.3979	100
50uM	1.699	100

Q-(C=O)-LD-CH$_2$-O-Ph

Caspase 3 w/ IE

FIGURE 28

FIGURE 29

IMPORTANT AMINO ACIDS